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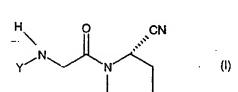
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(54) Title: 2-CYANOPYRROLIDINE DERIVATIVES AND THEIR USE AS MEDICAMENTS





(57) Abstract: The present invention relates to certain N-(substituted glycyl)-2-cyanopyrrolidines of formula (I) wherein Y is as defined herein, in free form or in acid addition salt form. Compounds of formula (I) inhibit DPP-IV (dipeptidyl-peptidase-IV) activity. They are therefore indicated for use as pharmaceuticals in inhibiting DPP-IV and in the treatment of conditions mediated by DPP-IV such as non-insulin-dependent diabetes mellitus, arthritis, obesity, osteoporosis and further conditions of impaired glucose tolerance.

'NTERNATIONAL SEARCH REPORT

rnational Application No PCT/EP 01/06595

A. CLASSIFICATION OF SUBJECT MATTER
IPC 7 C07D401/12 C07E C07D207/16 C07D403/12 C07D417/12 C07D409/12 A61K31/4439 A61K31/428 A61K31/427 A61K31/40 A61K31/506 A61P3/10 According to International Patent Classification (IPC) or to both national classification and IPC **B. FIELDS SEARCHED** Minimum documentation searched (classification system followed by classification symbols) IPC 7 C07D Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, CHEM ABS Data C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to claim No. Citation of document, with indication, where appropriate, of the relevant passages WO 98 19998 A (CIBA GEIGY AG; VILLHAUER 1 - 10EDWIN BERNARD (US)) 14 May 1998 (1998-05-14) claims; examples 1-66 US 6 011 155 A (VILLHAUER EDWIN BERNARD) Υ 1 - 104 January 2000 (2000-01-04) claims; examples T. E. HUGHES ET AL.: 1 - 10"(1-'''2-'(5-Cyanopyridin-2-yl!amino!acetyl!-2-cyano-(S)-pyrrolidine), a Slow-Binding Inhibitor of Dipeptidyl Peptidase IV" BIOCHEMISTRY, vol. 38, no. 36, 1999, pages 11597-11603, XP002184055 page 11597, abstract; page 11598, Chart 1 -/--Patent tamily members are listed in annex. Further documents are listed in the continuation of box C X Special categories of cited documents "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the 'A' document defining the general state of the art which is not considered to be of particular relevance. invention earlier document but published on or after the international . "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such docu-*O* document referring to an oral disclosure, use, exhibition or ments such combination being obvious to a person skilled document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 10/12/2001 26 November 2001 Authorized officer Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel (+31-70) 340-2040, Tx 31 651 epc nl. Hass, C

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'NTERNATIONAL SEARCH REPORT

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Category* Calation of document, with indication where appropriate, of the relevant passages Y BALKAN B ET AL: "INHIBITION OF DIPEPTIDYL PEPTIDASE IV WITH NWP—DPP728 INCREASES PLASMA GLP-1 (7-36 AMIDE) CONCENTRATIONS AND IMPROVES ORAL GLUCOSE TOLERANCE IN OBESE ZUCKER RATS" DIABETIOLOGIA, BERLIN, DE, vol. 42, no. 11, November 1999 (1999–11), pages 1324–1331, XP000921066 ISSN: 0012–186X page 1324, abstract; page 1325, figure 1 A EP 0 419 683 A (JAPAN TOBACCINC; 1-10 YOSHITOMI PHARMACEUTICAL (JP)) 3 April 1991 (1991–04-03) page 30, line 1 - line 12; claims			PCI/EP UI	./ 00595
PEPTIDASE IV WITH NVP-DPP728 INCREASES PLASMA GLP-1 (7-36 AMIDE) CONCENTRATIONS AND IMPROVES ORAL GLUCOSE TOLERANCE IN OBESE ZUCKER RATS" DIABETOLOGIA, BERLIN, DE, vol. 42, no. 11, November 1999 (1999-11), pages 1324-1331, XP000921066 ISSN: 0012-186X page 1324, abstract; page 1325, figure 1 A EP 0 419 683 A (JAPAN TOBACCO INC; YOSHITOMI PHARMACEUTICAL (JP)) 3 April 1991 (1991-04-03) page 30, line 1 - line 12; claims				Relevant to claim No.
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FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1-3, 7-10 (all partly)

Compounds of formula (I) wherein Y is a substituted 1,1-dimethylethyl group, and their pharmaceutical use

2. Claims: 1-3, 7-10 (all partly), 4-6

Compounds of formula (I) wherein Y is a substituted cyclohexyl group, and their pharmaceutical use

3. Claims: 1-3, 7-10 (all partly)

Compounds of formula (I) wherein Y is a substituted piperidinyl group, and their pharmaceutical use

4. Claims: 1-3, 7-10 (all partly)

Compounds of formula (I) wherein Y is a carbobicyclic system, and their pharmaceutical use

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